

This Page Is Inserted by IFW Operations
and is not a part of the Official Record

BEST AVAILABLE IMAGES

Defective images within this document are accurate representations of the original documents submitted by the applicant.

Defects in the images may include (but are not limited to):

- BLACK BORDERS
- TEXT CUT OFF AT TOP, BOTTOM OR SIDES
- FADED TEXT
- ILLEGIBLE TEXT
- SKEWED/SLANTED IMAGES
- COLORED PHOTOS
- BLACK OR VERY BLACK AND WHITE DARK PHOTOS
- GRAY SCALE DOCUMENTS

IMAGES ARE BEST AVAILABLE COPY.

**As rescanning documents *will not* correct images,
please do not report the images to the
Image Problem Mailbox.**

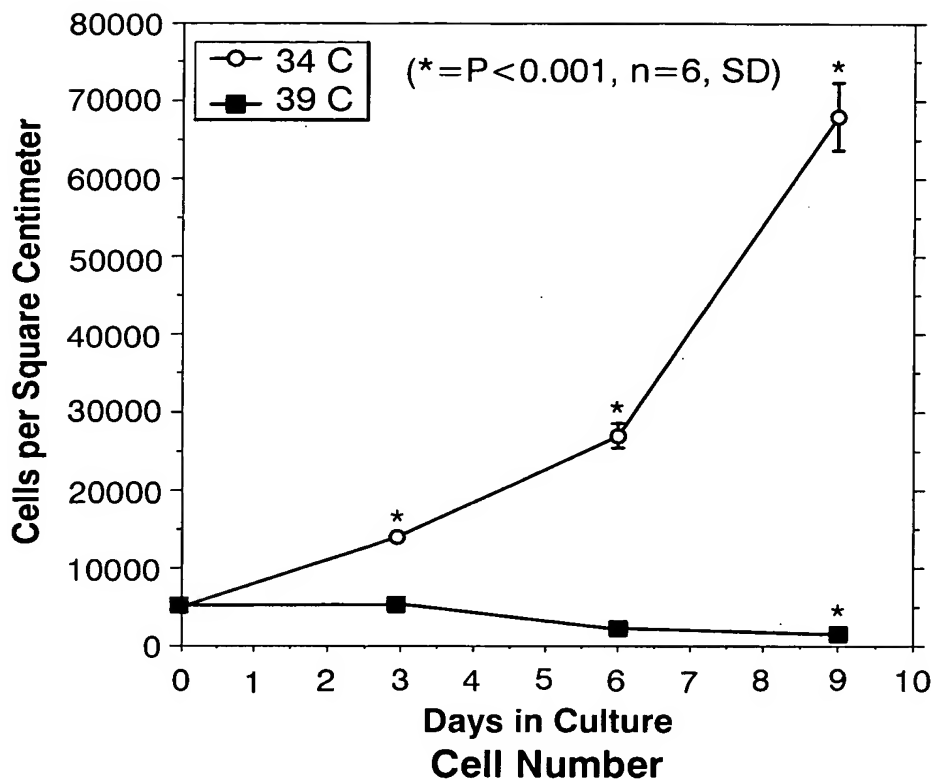
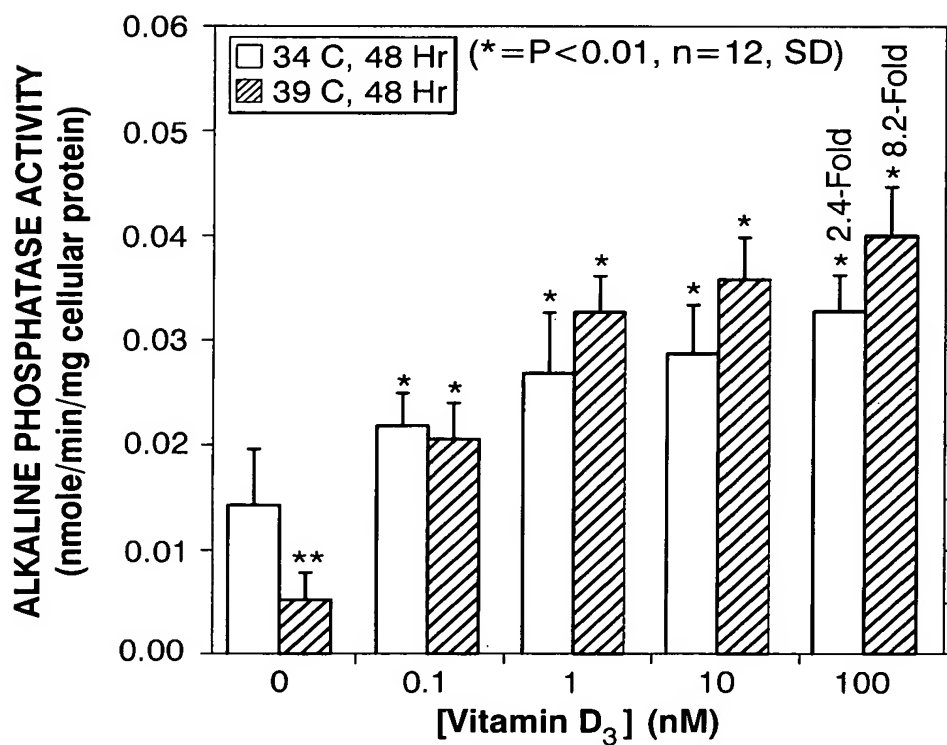
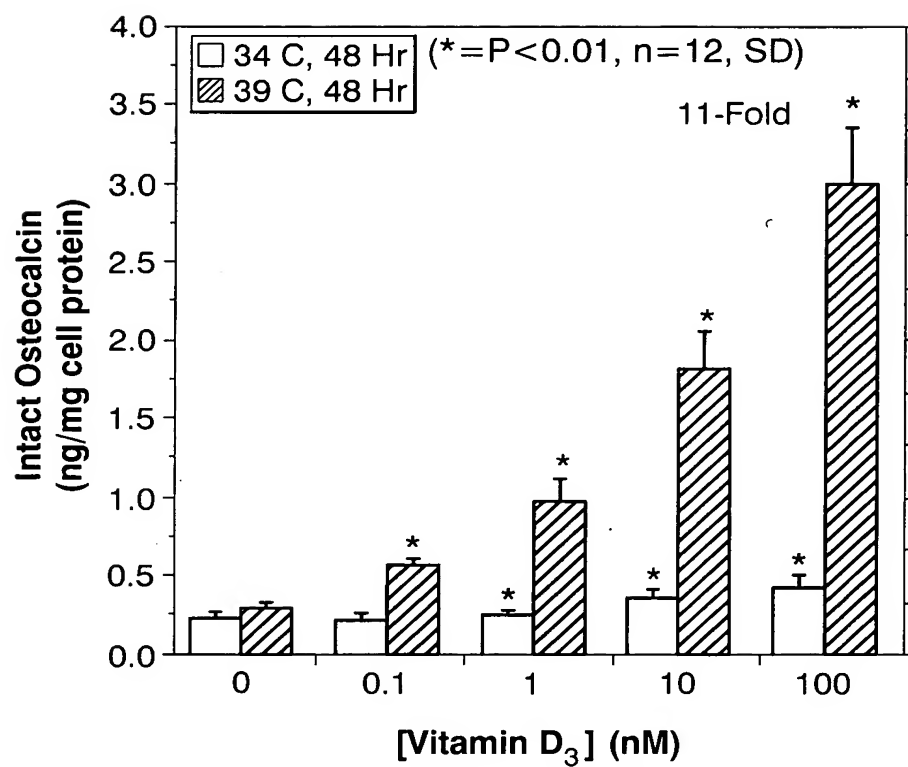


Figure 1A



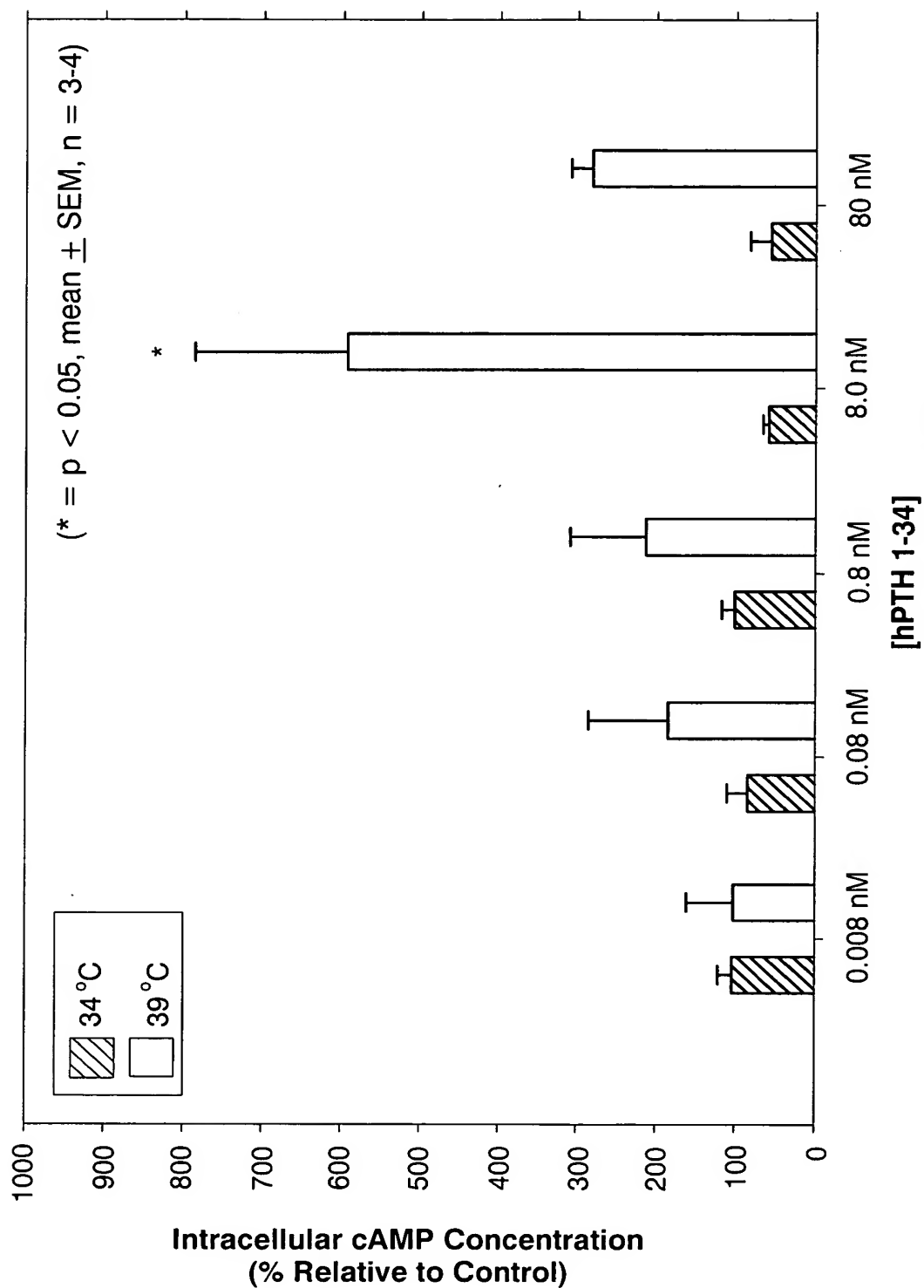
Alkaline Phosphatase Activity

Figure 1B



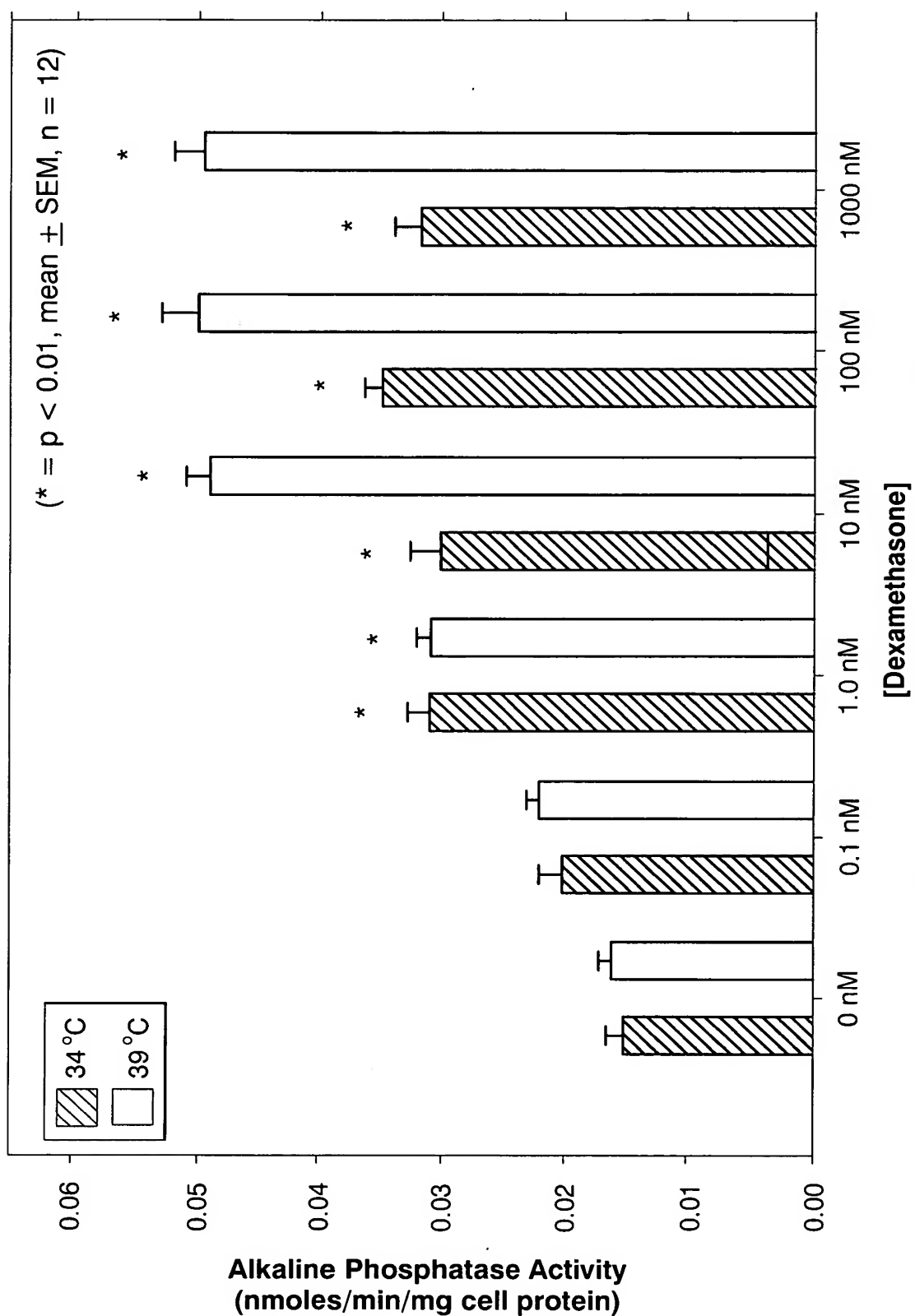
Osteocalcin Secretion

Figure 1C



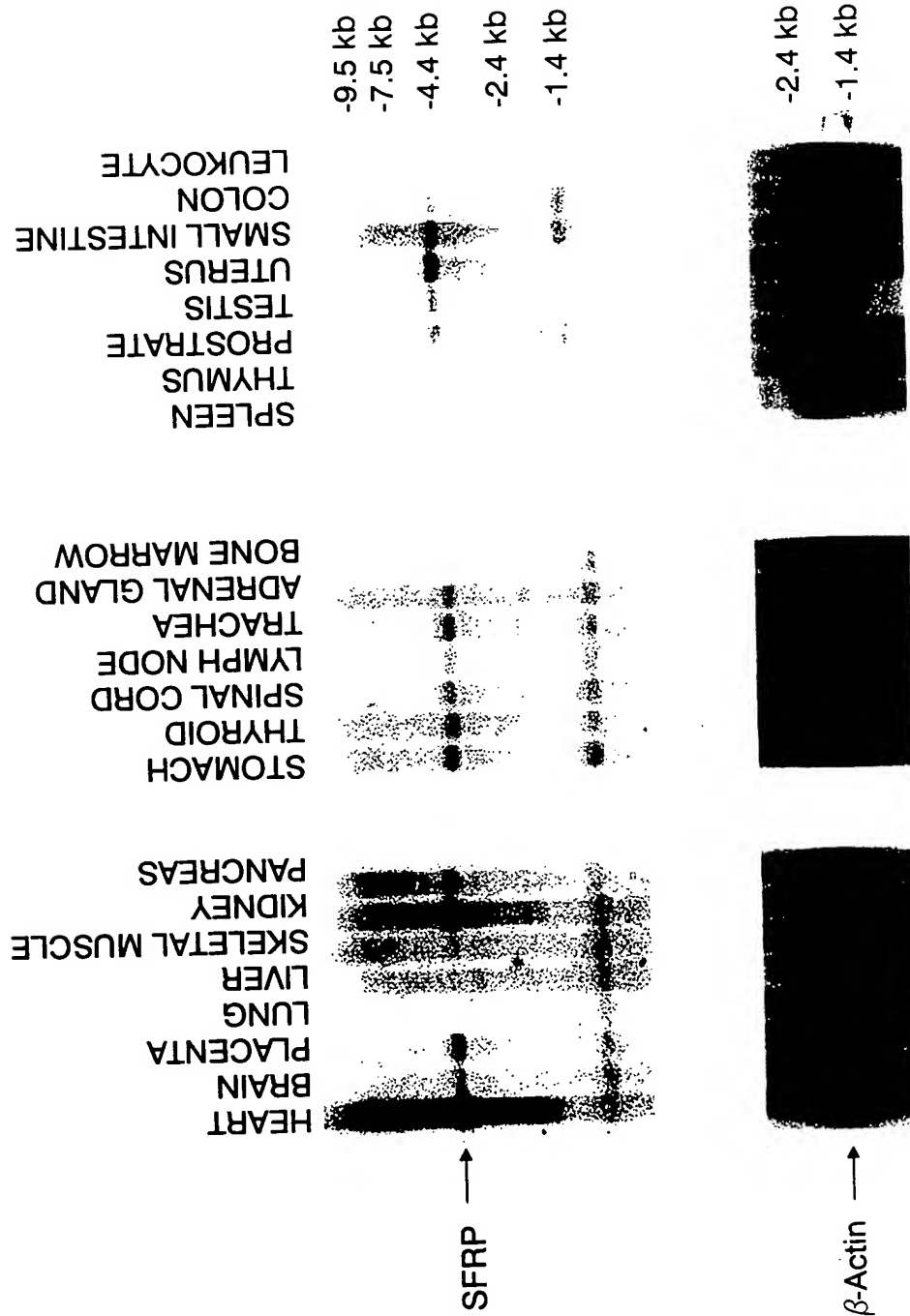
Regulation of Intracellular cAMP Levels
by Parathyroid Hormone in the HOB-01-C1-PS-09 Cells

Figure 2



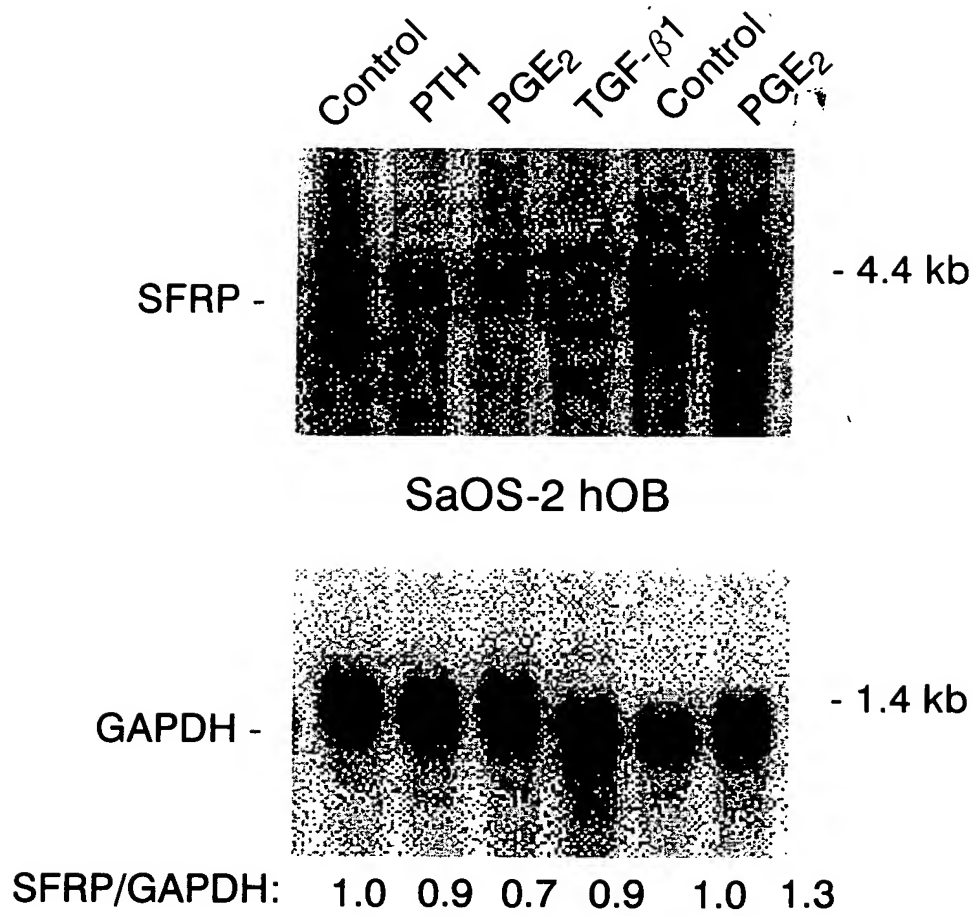
Regulation of Alkaline Phosphatase Activity
by Dexamethasone in the HOB-01-C1-PS-09 Cells

Figure 3



Osteogenic RADE Confirmation: SFRP

Figure 4



Osteogenic RADE Confirmation: SFRP

Figure 5

7/23

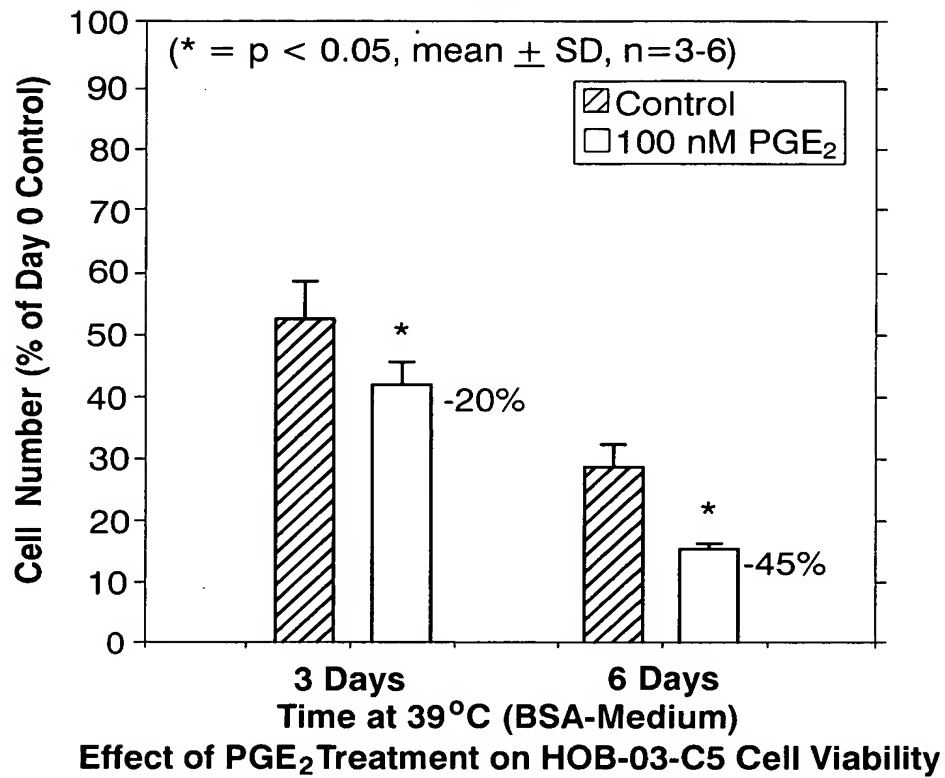


Figure 6A

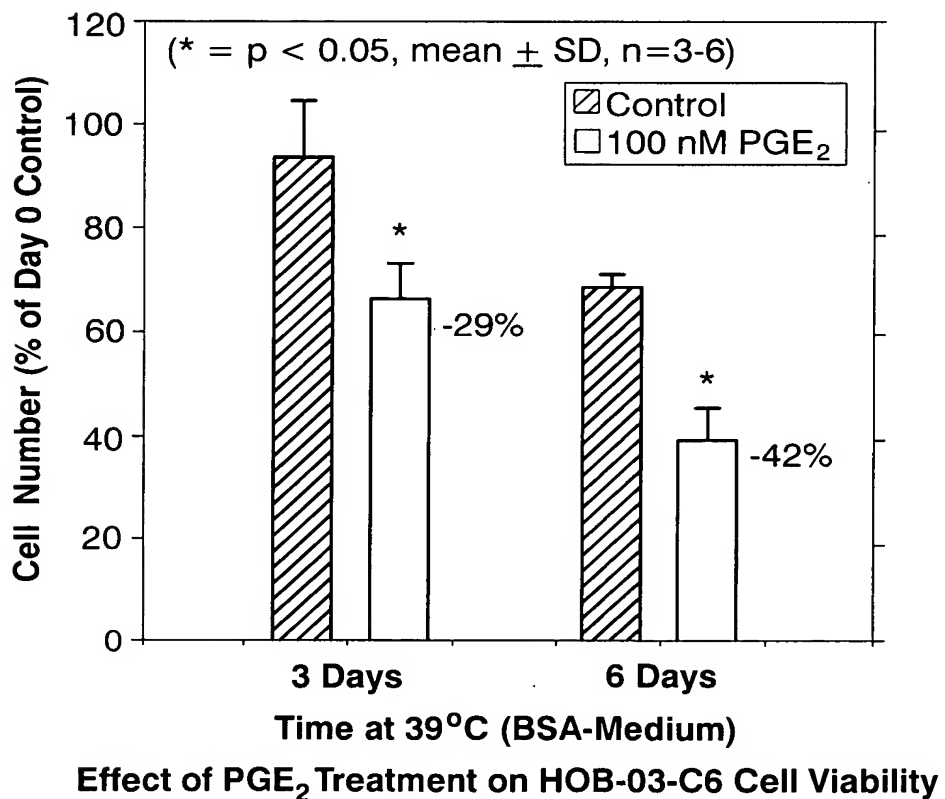
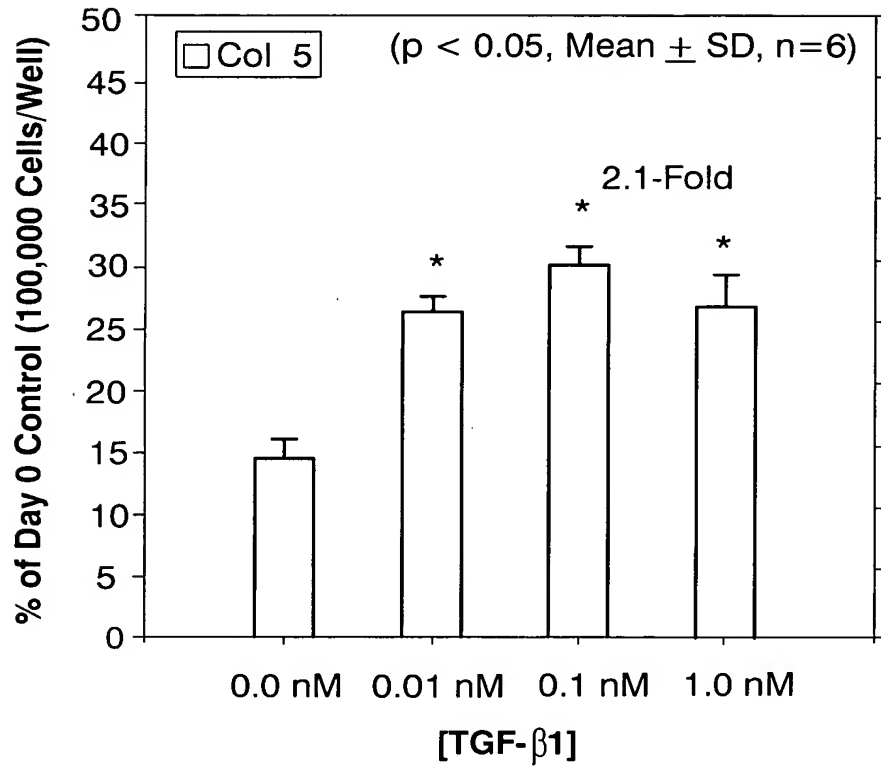
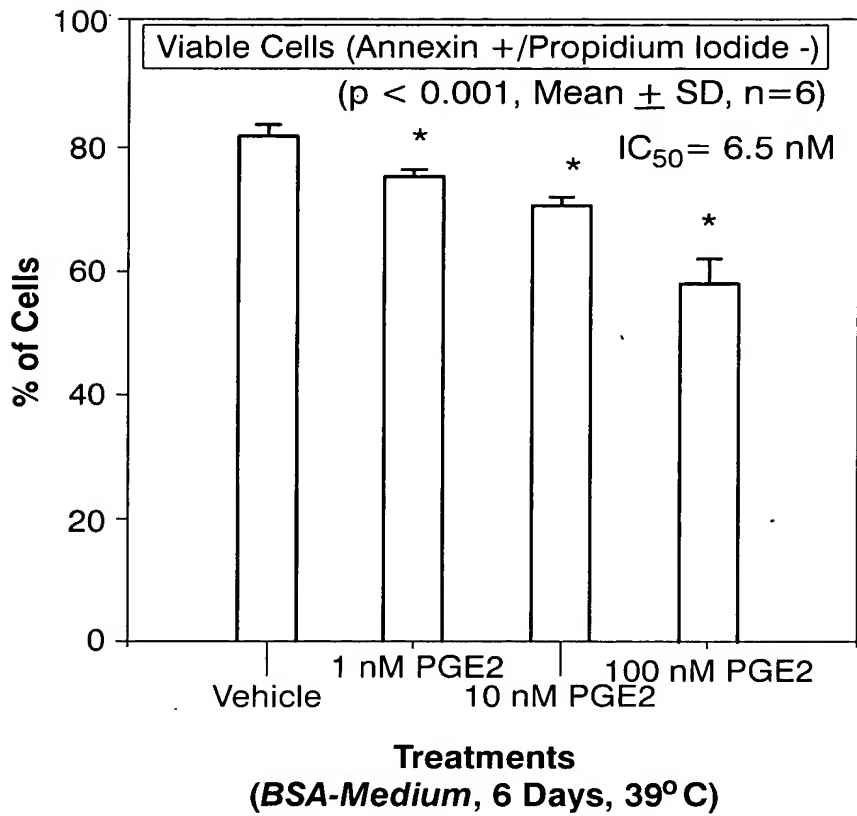


Figure 6B



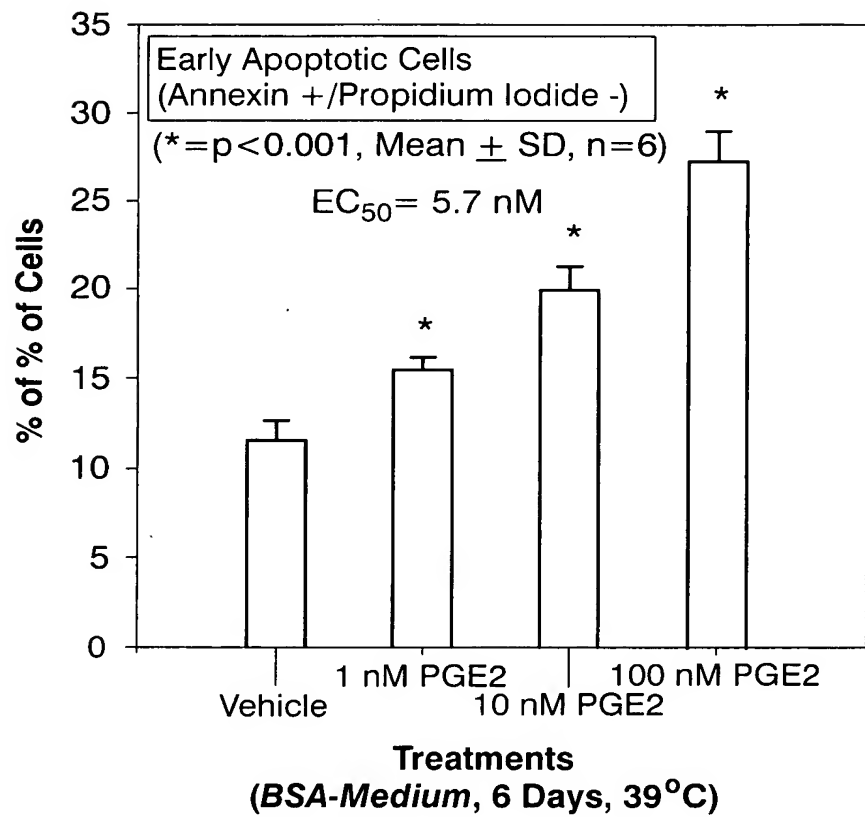
TGF- β 1 Treatment Increases HOB-01-C1 Cell Viability

Figure 6C



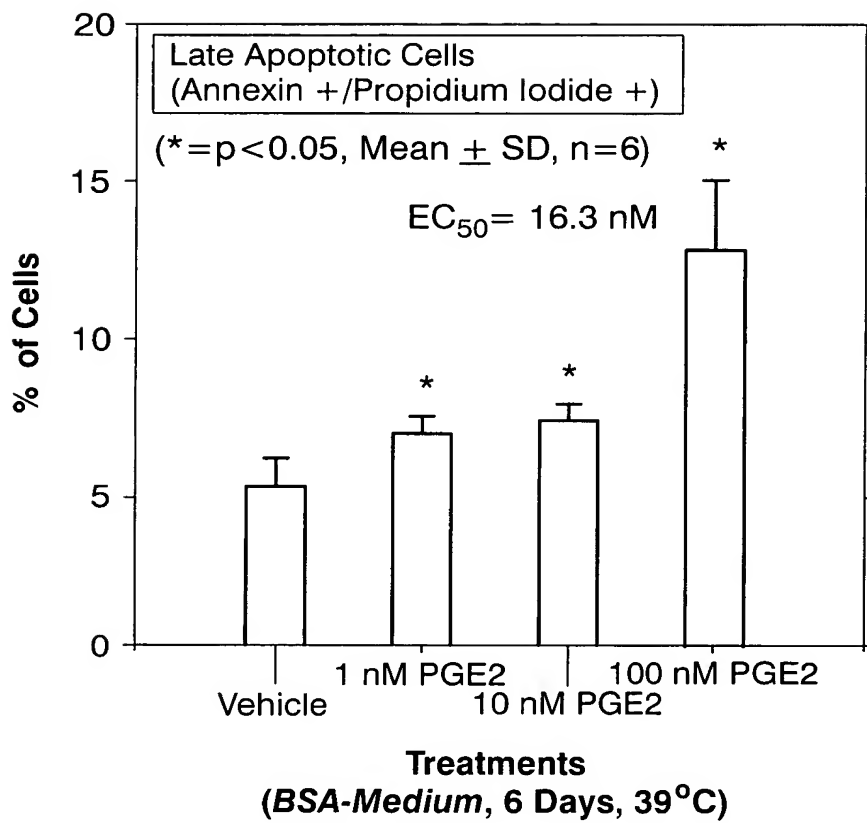
**Effect of PGE₂ on Proliferative-Stage
HOB-03-C5 Cell Apoptosis**

Figure 7A



Effect of PGE₂ on Proliferative-Stage
HOB-03-C5 Cell Apoptosis

Figure 7B



Effect of PGE₂ on Proliferative-Stage
HOB-03-C5 Cell Apoptosis

Figure 7C

12/23

**An Initiation Site-Directed Antisense Oligonucleotide for SARP-2
Reverses the Induction of Cell Death by PGE₂ in HOB-03-C5 Cells**

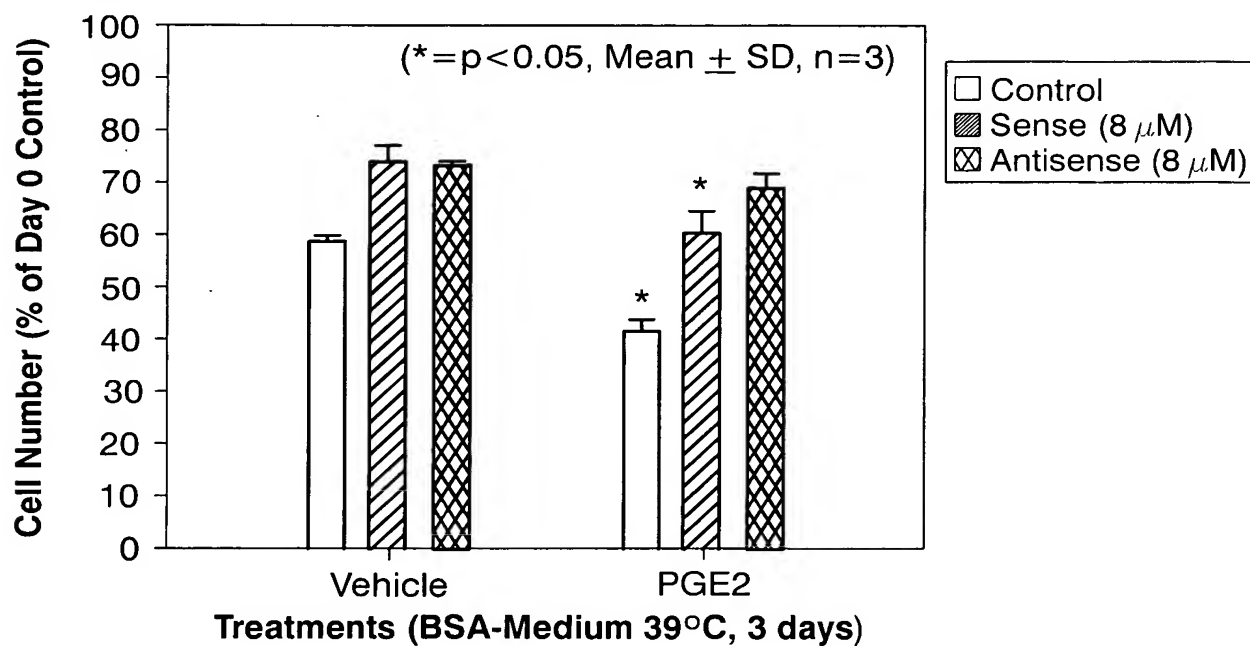


Figure 8A

**An Initiation Site-Directed Antisense Oligonucleotide for SARP-2
Reverses the Induction of Cell Death by PGE₂ in HOB-03-C6 Cells**

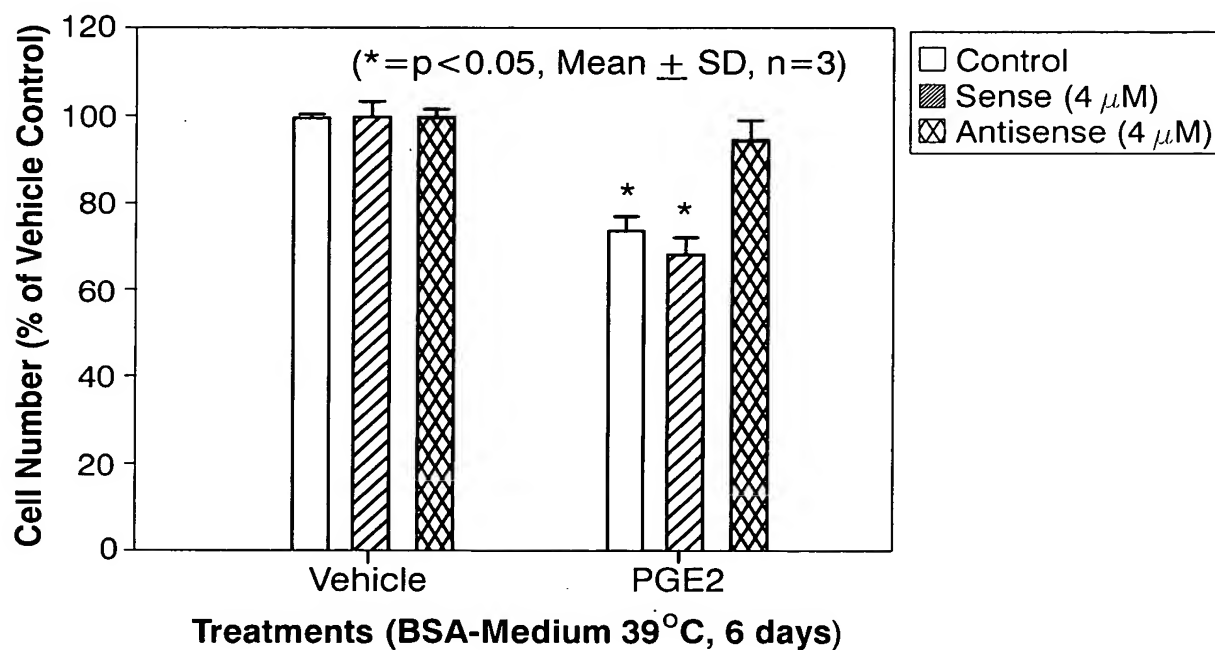
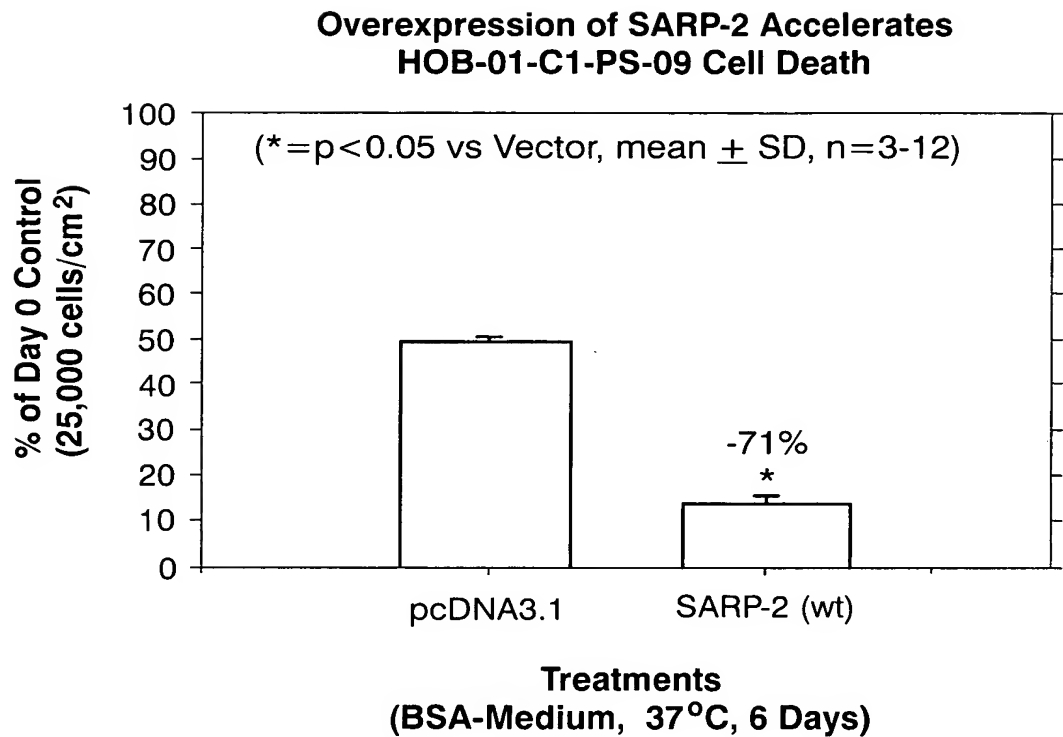


Figure 8B



SARP-2 Overexpression Accelerates HOB Cell Death
In Vitro Target Validation

Figure 9A

14/23

**Poly (A)+ RNA
Northern Blot**

**01-09
V S**

SARP -2



- 1.5 kb

GAPDH

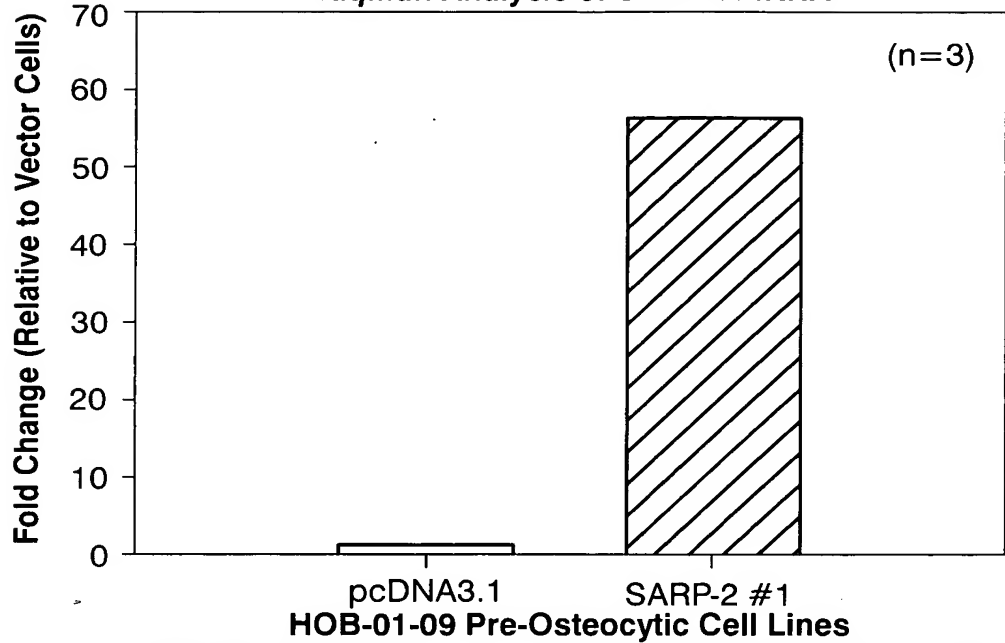


- 1.4 kb

Figure 9B

15/23

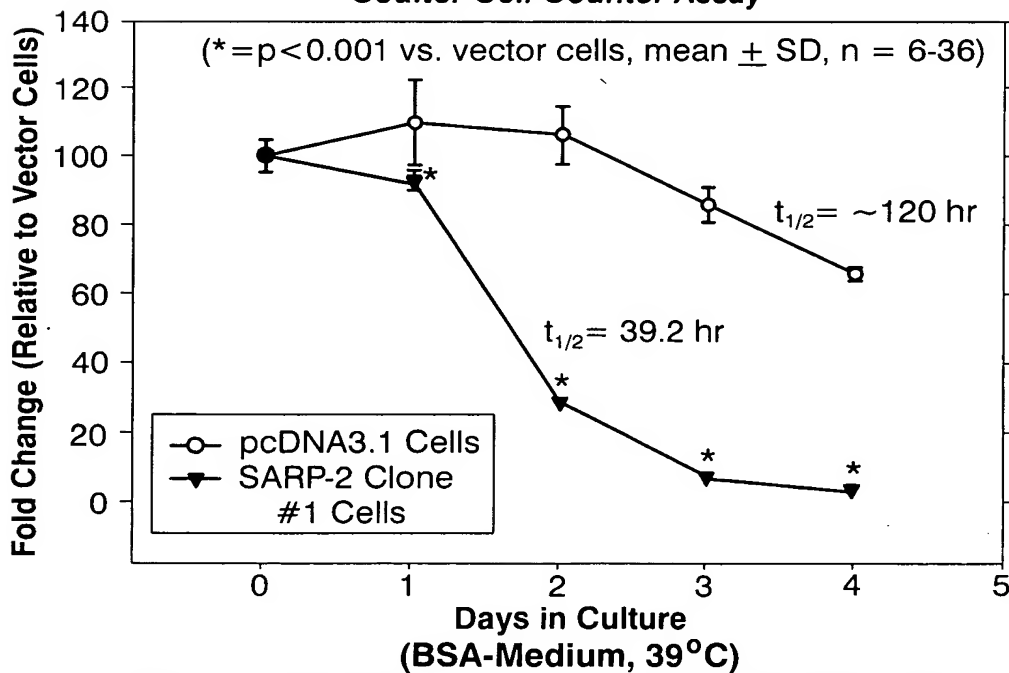
HOB-01-C1-PS-09 Cells
TaqMan Analysis of SARP-2 mRNA



SARP-2 Overexpression Accelerates HOB Cell Death
In Vitro Target Validation

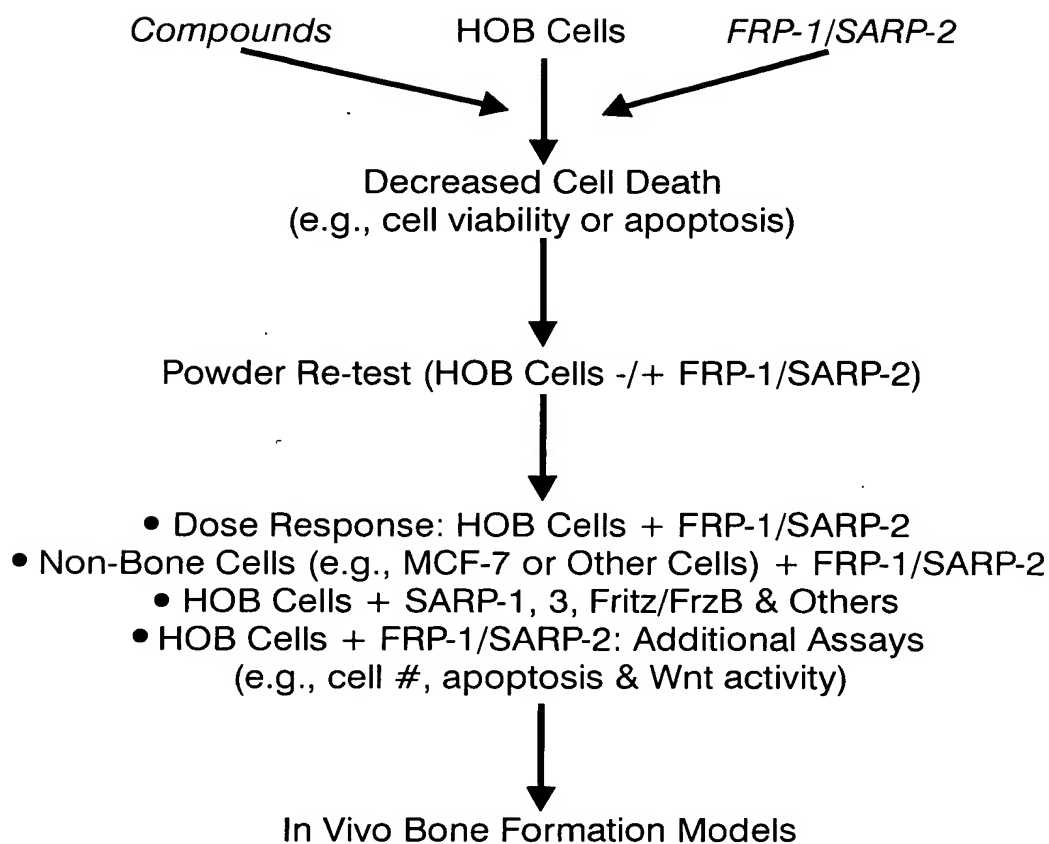
Figure 10A

HOB-01-C1-PS-09 Cells
Coulter Cell Counter Assay



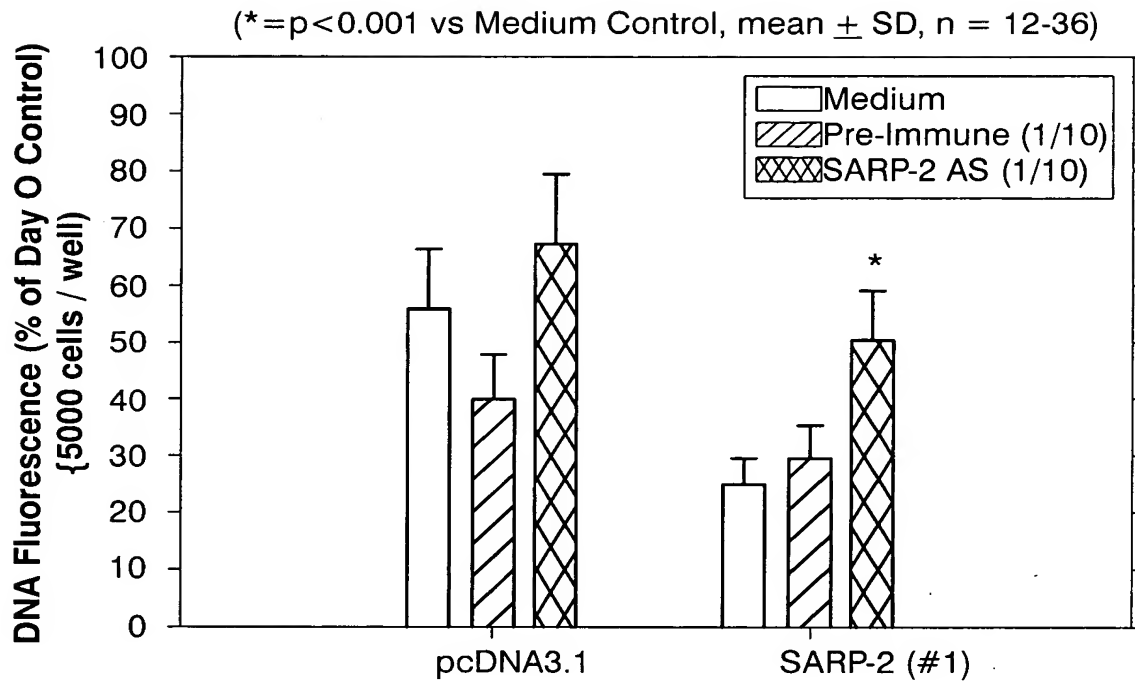
SARP-2 Overexpression Accelerates HOB Cell Death
In Vitro Target Validation

Figure 10B



**Use of Human FRP-1/SARP-2 and the HOB Cells
in a Screening Paradigm for an Anabolic Bone Agent**

Figure 11

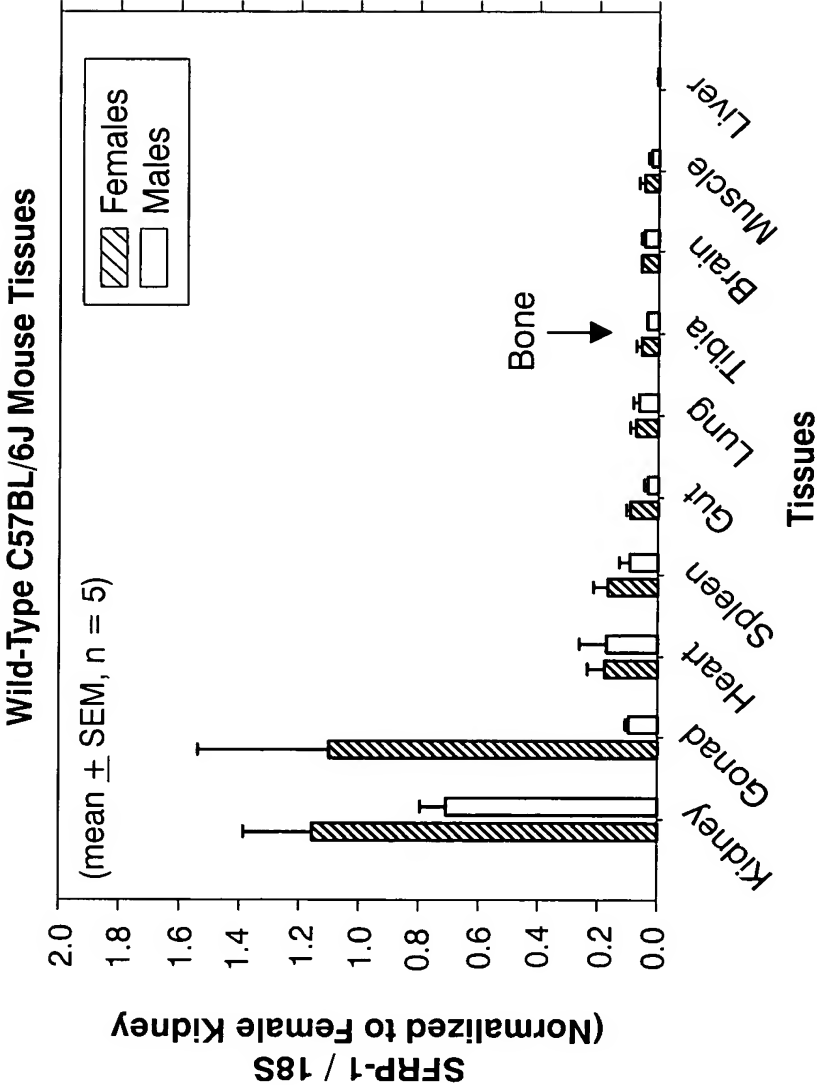


HOB-01-C1-PS-09 Cell Line

[BSA-Medium, 39°C, 3 Days]

SARP-2 Antisera Reverses SARP-2 Induced Cell Death
CyQuant HTS Assay

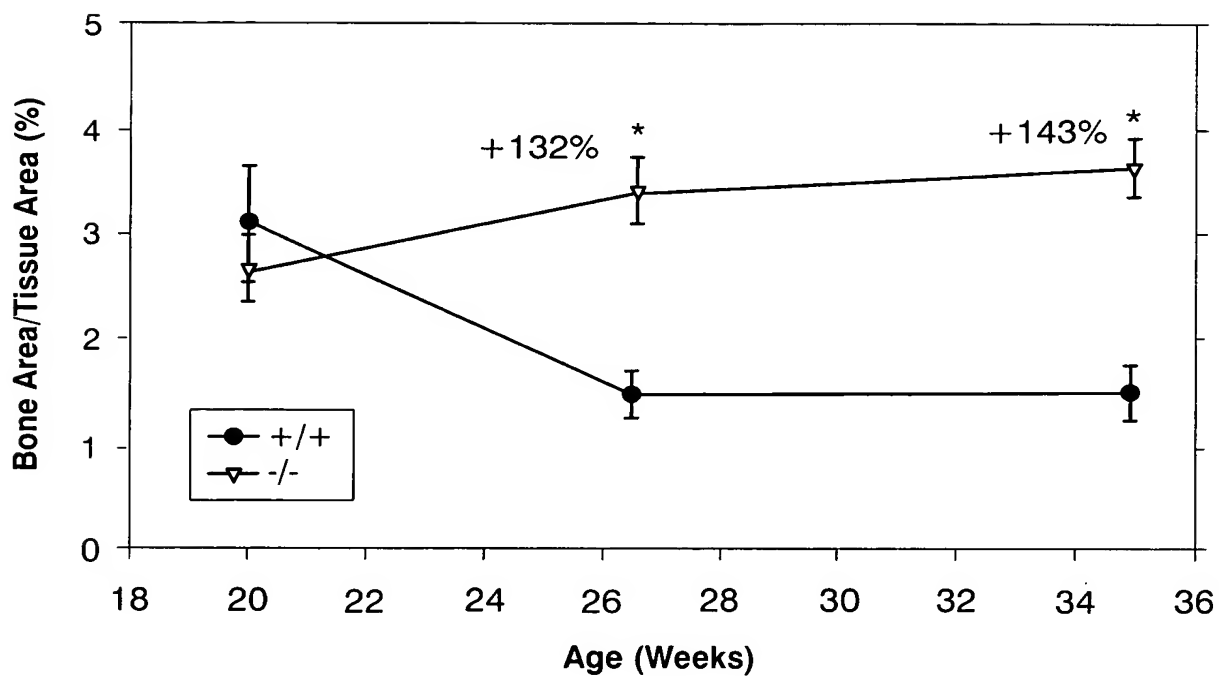
Figure 12



SFRP-1 mRNA is Highly Expressed in the Kidney & Ovary

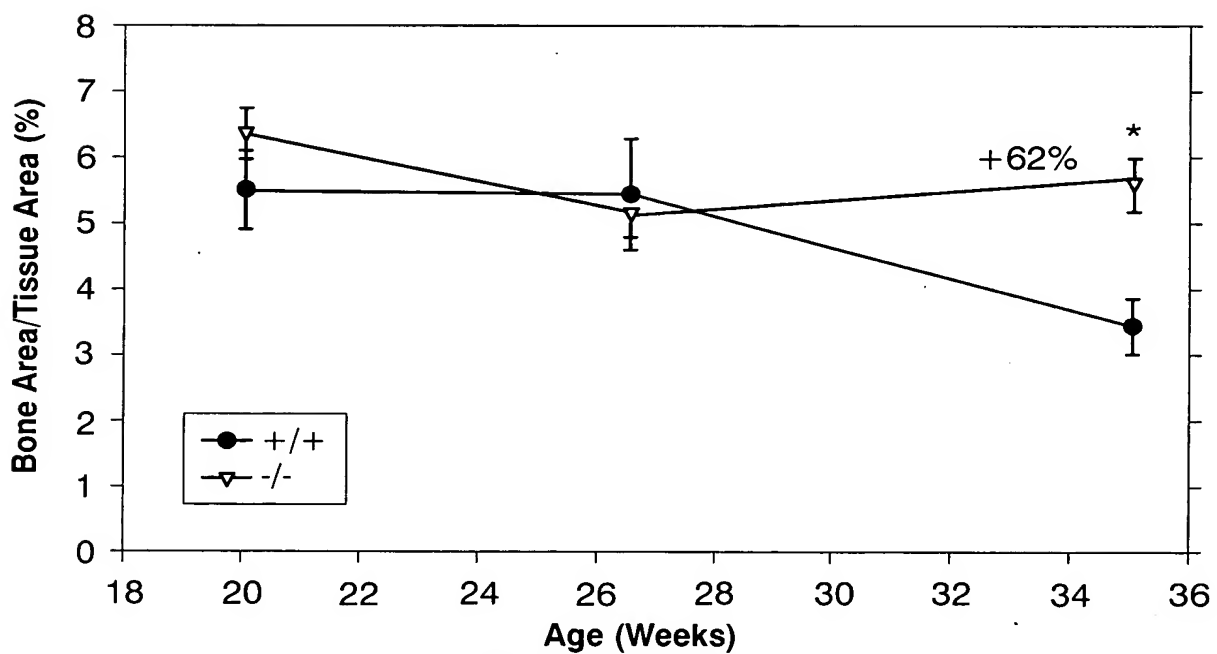
**Relative Tissue Distribution of Mouse SFRP-1
TaqMan Quantitative RT-PCR Analysis of Total RNA**

Figure 13



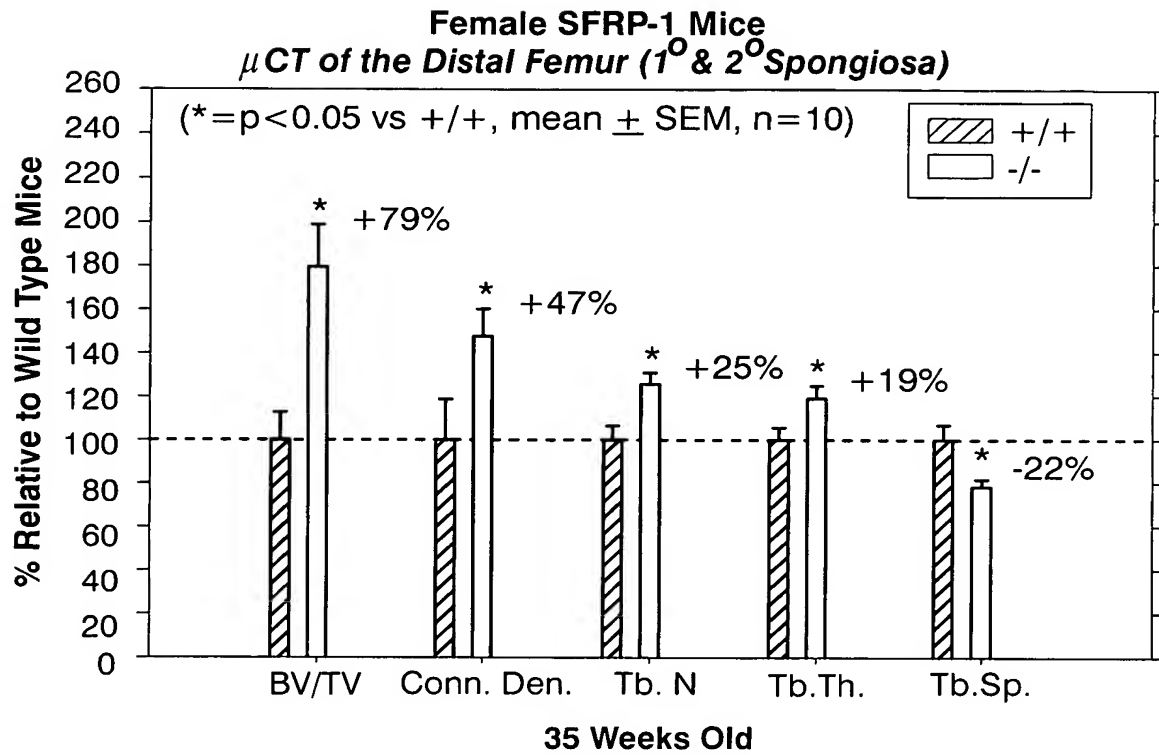
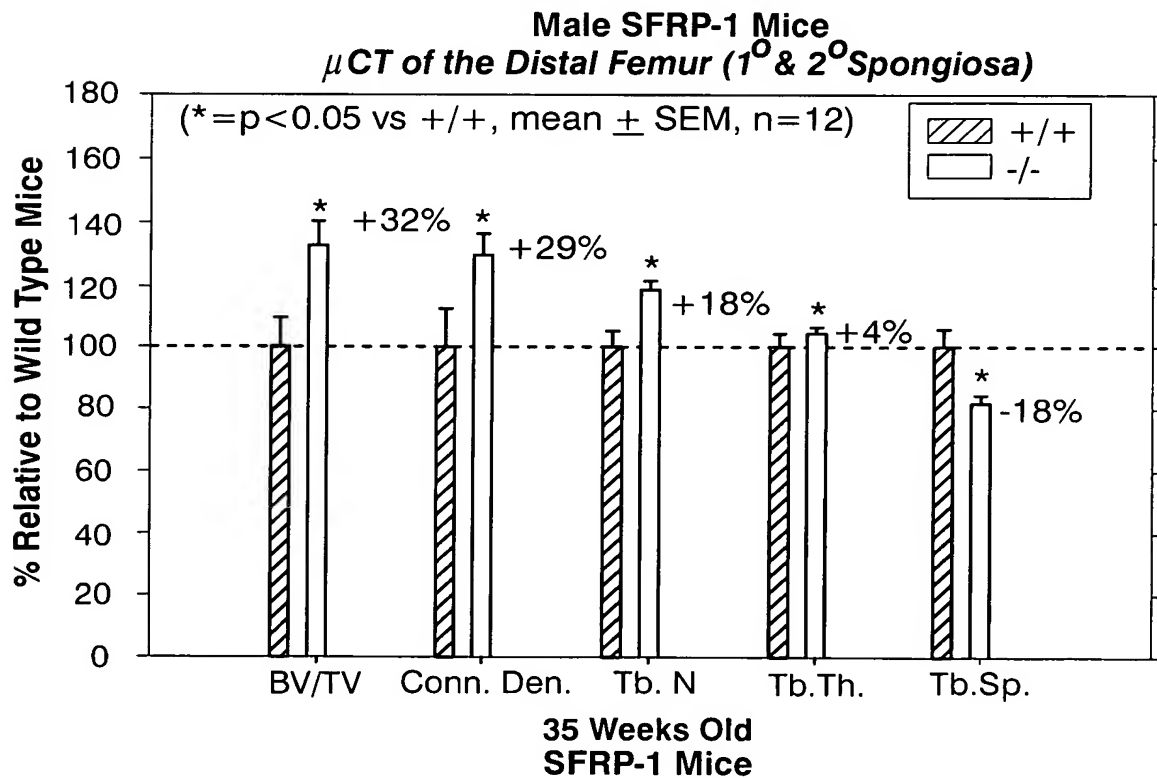
Female SFRP-1 Mice
Trabecular Bone Area

Figure 14A

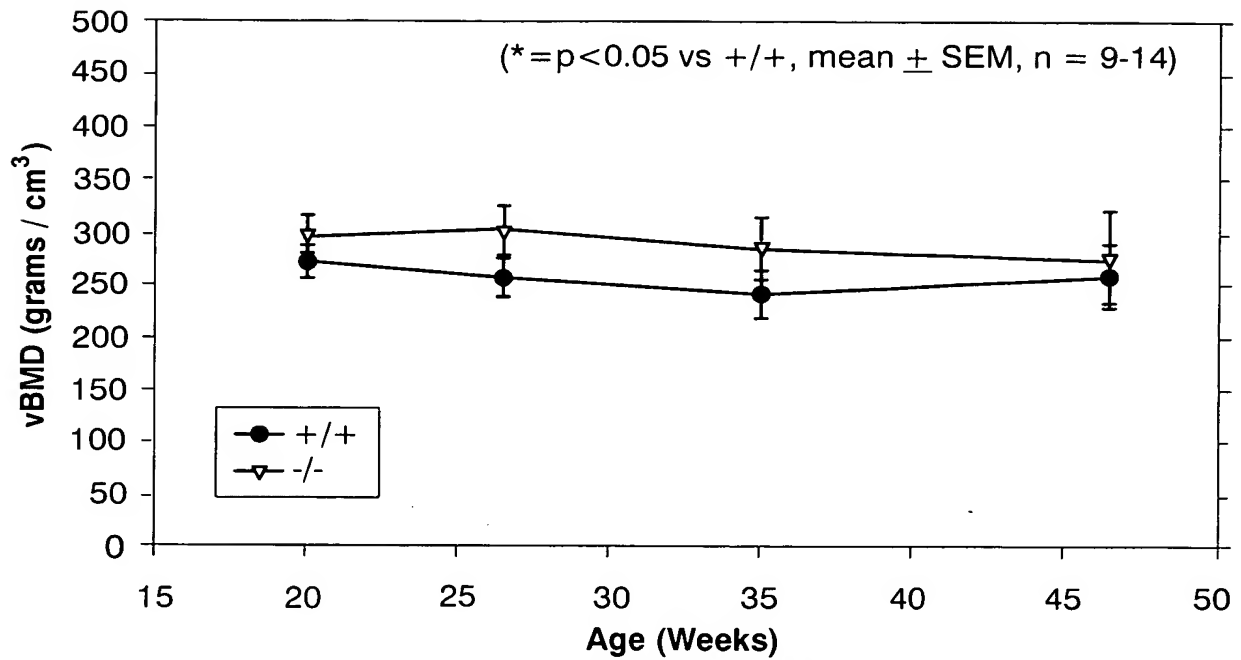


Male SFRP-1 Mice
Trabecular Bone Area

Figure 14B

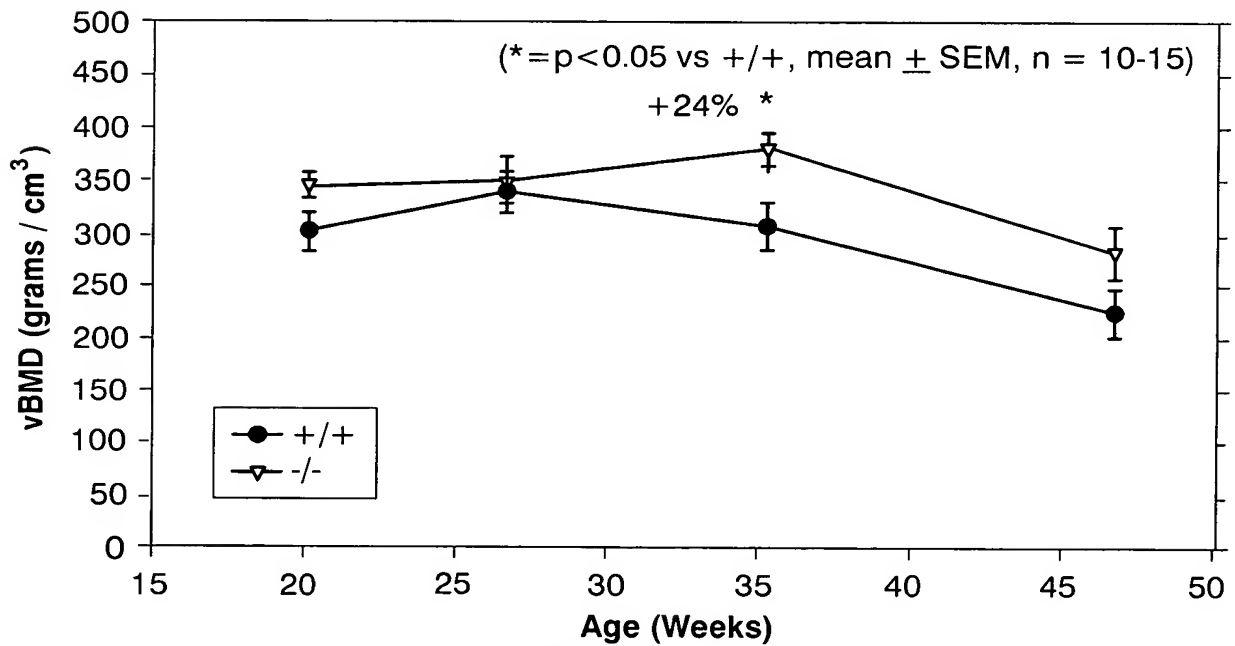
**Figure 15A****Figure 15B**

21/23



Female SFRP-1 Mice
Trabecular BMD of the Proximal Tibia (pQCT)
SFRP-1 KO Mice

Figure 16A



Male SFRP-1 Mice
Trabecular BMD of the Proximal Tibia (pQCT)
SFRP-1 KO Mice

Figure 16B

22/23

Female SFRP-1 Mice Bone Marrow Cells
Age 25-26 Weeks

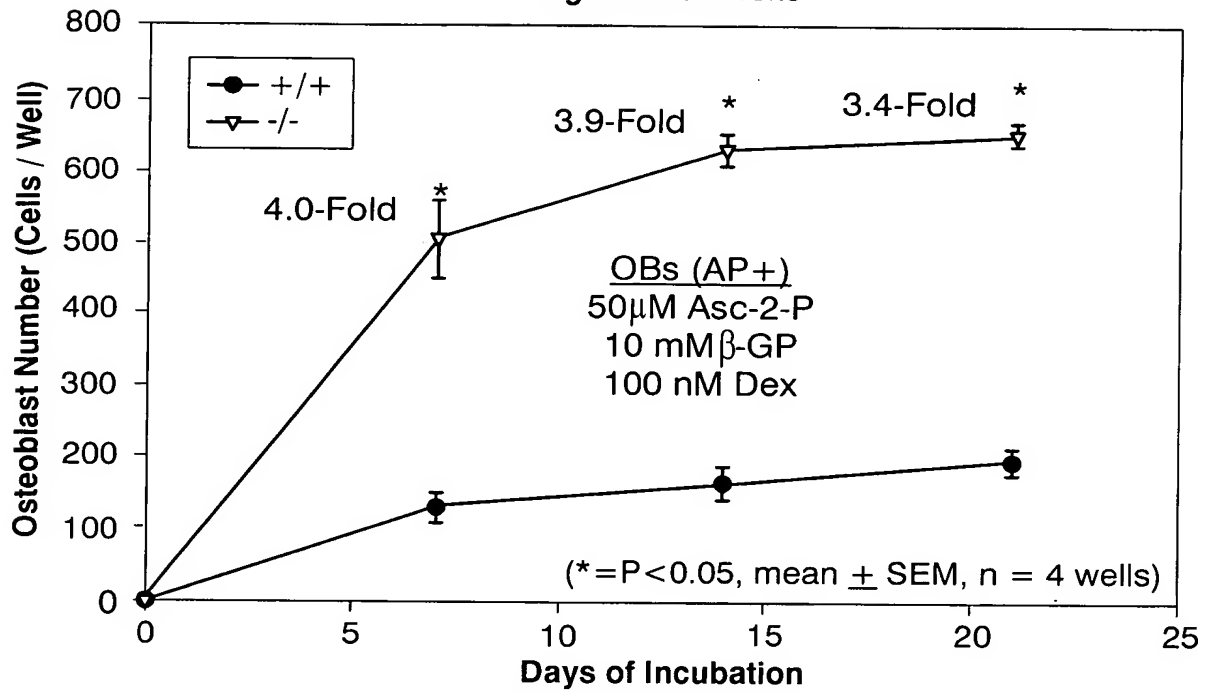


Figure 17A

Female SFRP-1 Mice Bone Marrow Cells
Age 25-26 Weeks

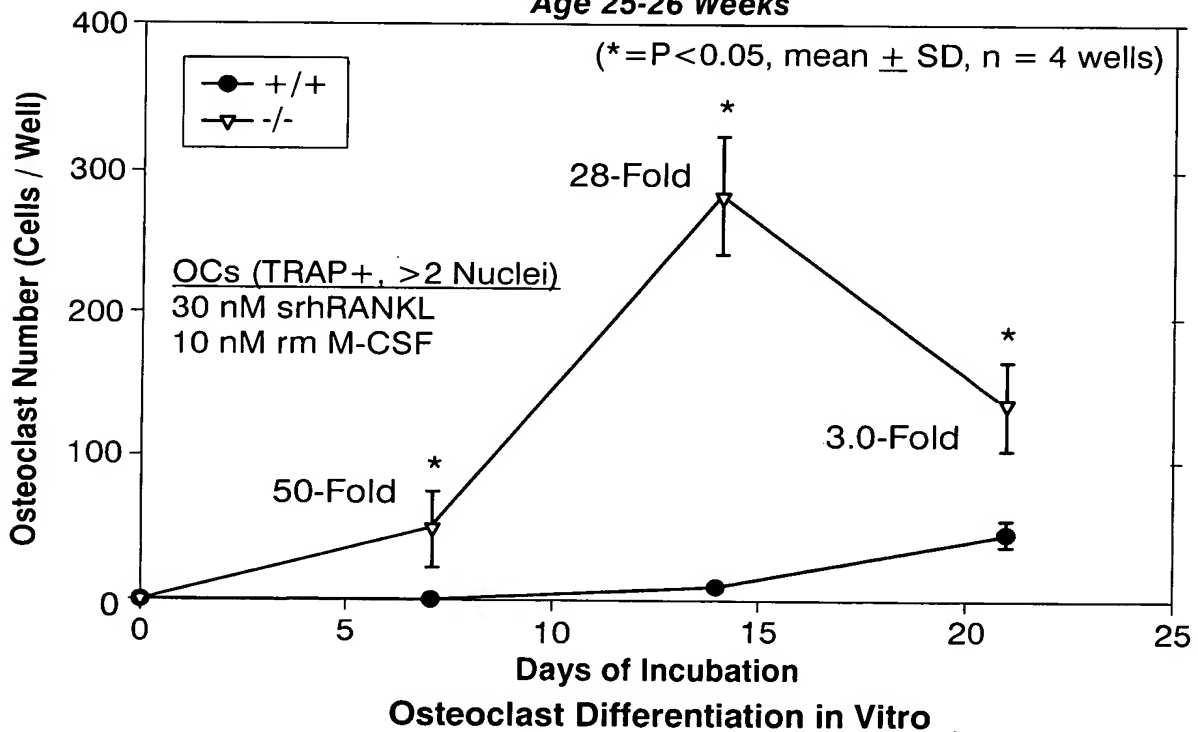
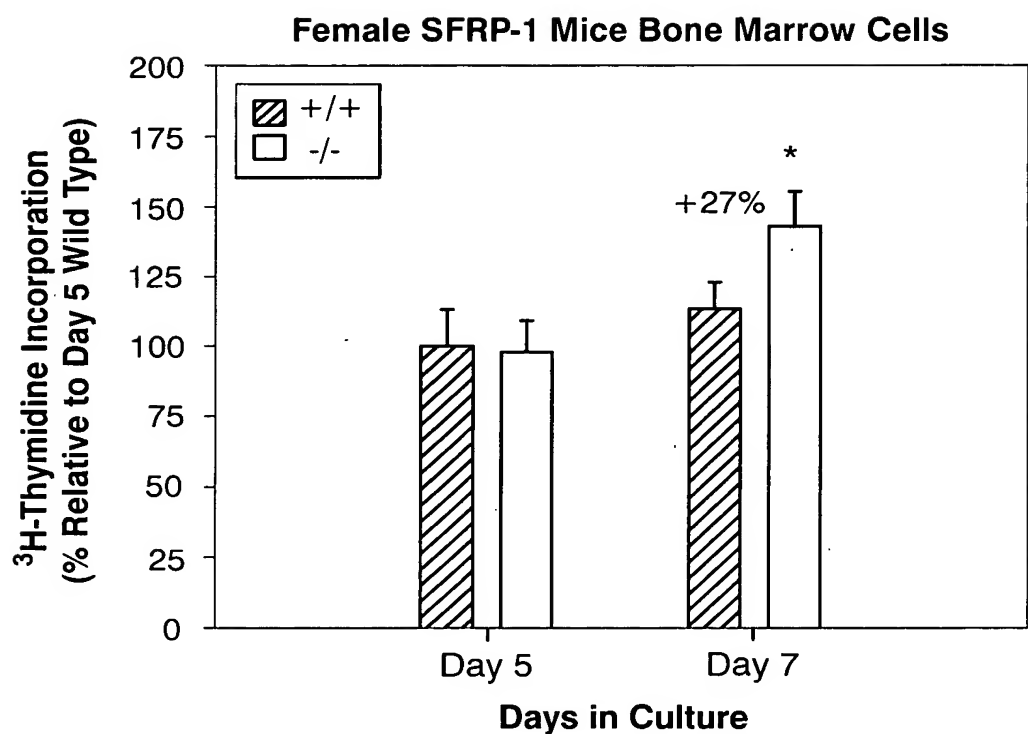
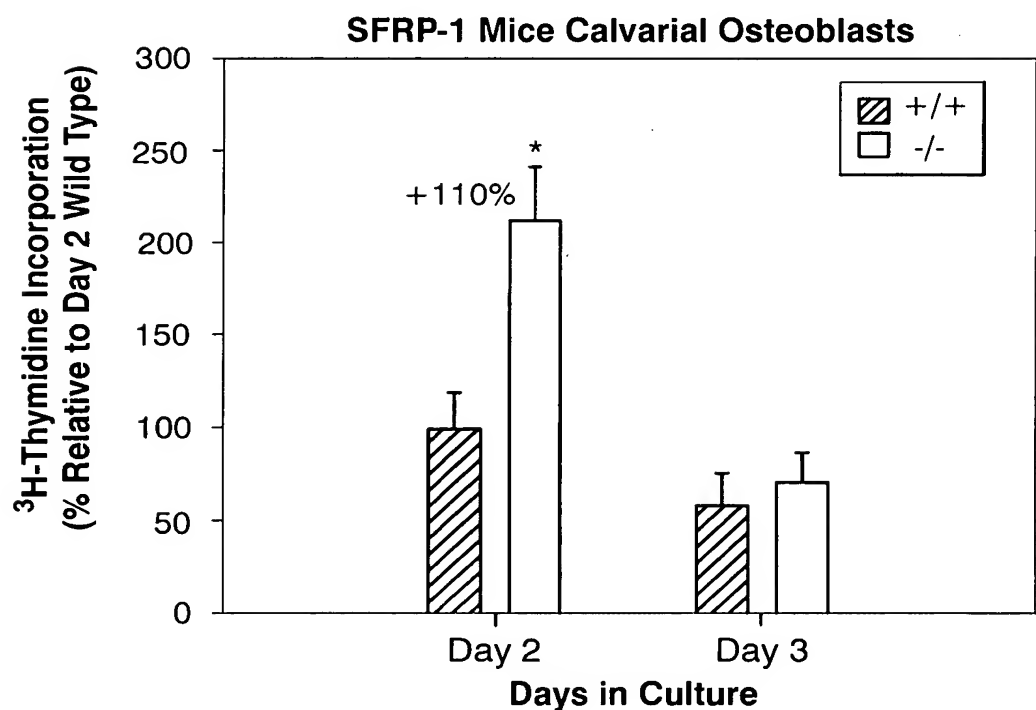


Figure 17B

**Figure 18A****Figure 18B**